

REMARKS

This Supplemental Preliminary Amendment is being filed to correct two, small and inadvertant errors introduced with the Preliminary Amendment filed May 6, 2002. The length and apparent complexity of this document is due solely to the required protocol for making amendments and not to any length or complexity of the amendments made.

In the Remarks portion of the May 6, 2002 Preliminary Amendment (at page 64 thereof), it was pointed out that the specification description contained an error with respect to the definition of certain of the rings. Specifically, it was noted that these rings were intended to be described as being 4-6 membered saturated heterocyclic rings instead of 5-6 membered saturated heterocyclic rings, for the reasons and with the support there described. Accordingly, a number of specification amendments were presented to make this correction. However, an error in the correction made at specification page 48 was recently noted, and it was further noted that this same error had been carried into new claim 39 presented in the Preliminary Amendment.

The purpose of this Supplemental Preliminary Amendment is to correct this ring definition at specification page 48 and in claim 39, and to make it consistent with the remainder of the specification and claims. Specifically:

- At page 48 of the specification, line14, R^{5a} was originally defined as “a 5- or 6-membered saturated heterocyclic group . . .” By the Preliminary Amendment of May 6, 2002, this was changed to read “a 4- or 6-membered saturated heterocyclic group . . .” However, as explained in the Remarks portion of the Preliminary Amendment, the intent was to have the definition refer to a 4-6 membered ring, and therefore the present specification

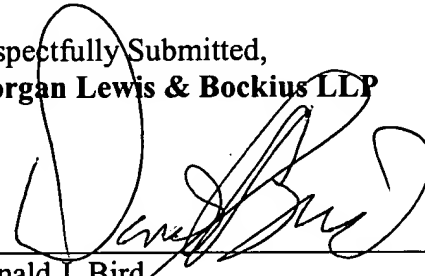
amendment corrects this definition of R^{5a} to read “a 4-, 5- or 6-membered saturated heterocyclic group.” However, to make this minor amendment, has been necessary to re-present the entire specification paragraph extending from page 47, line 26 through page 48, line 28, and to repeat this paragraph in the Appendix to show the amendment made. In this regard, the Examiner’s attention is called to the present Appendix at page 15, 5th line from the bottom, where the only specification change is shown by **bold underlined text**.

- Similarly, in claim 39 as presented in the Preliminary Amendment of May 6, 2002, the definition of R^{5a} was erroneously presented as “a 4- or 6-membered saturated heterocyclic group . . .” See Preliminary Amendment at page 48, line 14-15. This is corrected by the present amendment to read “a 4-, 5- or 6-membered saturated heterocyclic group.” In this regard, the Examiner’s attention is called to the present Appendix at page 24, line 4, where the only change to claim 39 is shown by **bold underlined text**.

Conclusion

For the above reasons, and as explained in the Preliminary Amendment, no new matter is introduced by the above amendments. In that they simply correct an inadvertant error in the Preliminary Amendment, entry thereof is believed to be in order and is respectfully requested.

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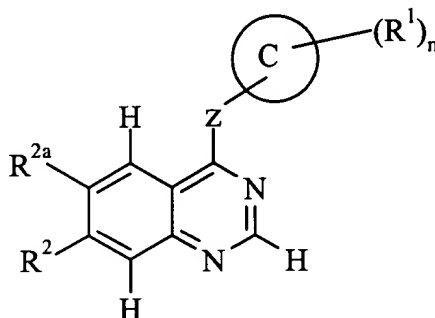
APPENDIX
VERSION WITH MARKINGS TO SHOW CHANGES

IN THE SPECIFICATION:

The specification has been amended as shown below, wherein additions are indicated by text in **bold underline**, and deletions are indicated by text **[in bold between brackets]**:

The paragraph extending from page 47, line 26 to page 48, line 28 has been amended as follows:

In another aspect of the present invention there is provided the use of compounds of the formula Ia:



(Ia)

[wherein:

ring C, R^1 , R^2 , n and Z are as defined hereinbefore with the provisos that R^2 is not hydrogen and that Z is not CH_2 or a direct bond; and

R^{2a} represents hydrogen, halogeno, C_{1-3} alkyl, trifluoromethyl, C_{1-3} alkoxy, C_{1-3} alkylsulphanyl, $-NR^{3a}R^{4a}$ (wherein R^{3a} and R^{4a} , which may be the same or different, each represents hydrogen or C_{1-3} alkyl), or $R^{5a}(CH_2)_{za}X^{1a}$ (wherein R^{5a} is a 4-, 5- or 6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which heterocyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C_{1-4} cyanoalkyl, C_{1-4} alkyl, C_{1-4} hydroxyalkyl, C_{1-4} alkoxy, C_{1-4} alkoxy C_{1-4} alkyl, C_{1-4} alkylsulphonyl C_{1-4} alkyl, C_{1-4} alkoxycarbonyl, C_{1-4} aminoalkyl, C_{1-4} alkylamino, di(C_{1-4}

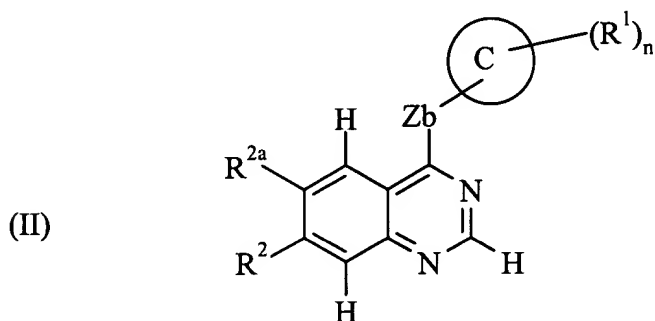
$_4$ alkyl)amino, C_{1-4} alkylamino C_{1-4} alkyl, di(C_{1-4} alkyl)amino C_{1-4} alkyl, C_{1-4} alkylamino C_{1-4} alkoxy, di(C_{1-4} alkyl)amino C_{1-4} alkoxy and a group $-(-O-)_f(C_{1-4}alkyl)_g ringD$ (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C_{1-4} alkyl), z_a is an integer from 0 to 4 and X^{1a} represents a direct bond, -O-, -CH₂-, -S-, -SO-, -SO₂-, -NR^{6a}C(O)-, -C(O)NR^{7a}-, -SO₂NR^{8a}-, -NR^{9a}SO₂- or -NR^{10a}- (wherein R^{6a}, R^{7a}, R^{8a}, R^{9a} and R^{10a} each independently represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl));

and salts thereof, and prodrugs thereof for example esters and amides, in the manufacture of a medicament for use in the production of an antiangiogenic and/or vascular permeability reducing effect in warm-blooded animals such as humans.

IN THE CLAIMS:

Claim 39 has been amended as follows:

39. (Amended) A compound of the formula II:



wherein:

ring C is an 8, 9, 10, 12 or 13-membered bicyclic or tricyclic moiety which moiety may be saturated or unsaturated, which may be aromatic or non-aromatic, and which optionally may contain 1-3 heteroatoms selected independently from O, N and S;

Z_b is -O- or -S-;

n is an integer from 0 to 5;

R^2 represents hydrogen, hydroxy, halogeno, cyano, nitro, trifluoromethyl, C_{1-3} alkyl, C_{1-3} alkoxy, C_{1-3} alkylsulphanyl, $-NR^3R^4$ (wherein R^3 and R^4 , which may be the same or different, each represents hydrogen or C_{1-3} alkyl),

or R^2 represents a group R^5X^1- , wherein X^1 represents a direct bond, $-O-$, $-CH_2-$, $-OC(O)-$, $-C(O)-$, $-S-$, $-SO-$, $-SO_2-$, $-NR^6C(O)-$, $-C(O)NR^7-$, $-SO_2NR^8-$, $-NR^9SO_2-$ or $-NR^{10}-$ (wherein R^6 , R^7 , R^8 , R^9 and R^{10} each independently represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl), and R^5 is selected from one of the following twenty-two groups:

- 1) hydrogen, oxiranyl C_{1-4} alkyl or C_{1-5} alkyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, chloro, bromo and amino;
- 2) C_{1-5} alkyl $X^2C(O)R^{11}$ (wherein X^2 represents $-O-$ or $-NR^{12}-$ (in which R^{12} represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl) and R^{11} represents C_{1-3} alkyl, $-NR^{13}R^{14}$ or $-OR^{15}$ (wherein R^{13} , R^{14} and R^{15} which may be the same or different each represents hydrogen, C_{1-5} alkyl or C_{1-3} alkoxy C_{2-3} alkyl));
- 3) C_{1-5} alkyl X^3R^{16} (wherein X^3 represents $-O-$, $-S-$, $-SO-$, $-SO_2-$, $-OC(O)-$, $-NR^{17}C(O)-$, $-C(O)NR^{18}-$, $-SO_2NR^{19}-$, $-NR^{20}SO_2-$ or $-NR^{21}-$ (wherein R^{17} , R^{18} , R^{19} , R^{20} and R^{21} each independently represents hydrogen, C_{1-3} alkyl or C_{1-3} alkoxy C_{2-3} alkyl) and R^{16} represents hydrogen, C_{1-3} alkyl, cyclopentyl, cyclohexyl or a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which C_{1-3} alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and C_{1-4} alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C_{1-4} cyanoalkyl, C_{1-4} alkyl, C_{1-4} hydroxyalkyl, C_{1-4} alkoxy, C_{1-4} alkoxy C_{1-4} alkyl, C_{1-4} alkylsulphonyl C_{1-4} alkyl, C_{1-4} alkoxycarbonyl, C_{1-4} aminoalkyl, C_{1-4} alkylamino, di(C_{1-4} alkyl)amino, C_{1-4} alkylamino C_{1-4} alkyl, di(C_{1-4} alkyl)amino C_{1-4} alkyl, C_{1-4} alkylamino C_{1-4} alkoxy, di(C_{1-4} alkyl)amino C_{1-4} alkoxy and a group $-(O-)_f(C_{1-4}alkyl)_g$ ringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C_{1-4} alkyl));

- 4) $C_{1-5}alkylX^4C_{1-5}alkylX^5R^{22}$ (wherein X^4 and X^5 which may be the same or different are each -O-, -S-, -SO-, -SO₂-, -NR²³C(O)-, -C(O)NR²⁴-, -SO₂NR²⁵-, -NR²⁶SO₂- or -NR²⁷- (wherein R²³, R²⁴, R²⁵, R²⁶ and R²⁷ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²² represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl);
- 5) R²⁸ (wherein R²⁸ is a 4-6-membered saturated heterocyclic group (linked via carbon or nitrogen) with 1-2 heteroatoms, selected independently from O, S and N, which heterocyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₄cyanoalkyl, C₁₋₄alkyl, C₁₋₄hydroxyalkyl, C₁₋₄alkoxy, C₁₋₄alkoxyC₁₋₄alkyl, C₁₋₄alkylsulphonylC₁₋₄alkyl, C₁₋₄alkoxycarbonyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, C₁₋₄alkylaminoC₁₋₄alkoxy, di(C₁₋₄alkyl)aminoC₁₋₄alkoxy and a group $-(O)_f(C_{1-4}alkyl)_gringD$ (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl));
- 6) $C_{1-5}alkylR^{28}$ (wherein R²⁸ is as defined herein);
- 7) $C_{2-5}alkenylR^{28}$ (wherein R²⁸ is as defined herein);
- 8) $C_{2-5}alkynylR^{28}$ (wherein R²⁸ is as defined herein);
- 9) R²⁹ (wherein R²⁹ represents a pyridone group, a phenyl group or a 5-6-membered aromatic heterocyclic group (linked via carbon or nitrogen) with 1-3 heteroatoms selected from O, N and S, which pyridone, phenyl or aromatic heterocyclic group may carry up to 5 substituents selected from hydroxy, halogeno, amino, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄hydroxyalkyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, C₁₋₄hydroxyalkoxy, carboxy, trifluoromethyl, cyano, -C(O)NR³⁰R³¹, -NR³²C(O)R³³ (wherein R³⁰, R³¹, R³² and R³³, which may be the same or different, each represents hydrogen, C₁₋₄alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and a group $-(O)_f(C_{1-4}alkyl)_gringD$ (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl));
- 10) $C_{1-5}alkylR^{29}$ (wherein R²⁹ is as defined herein);
- 11) $C_{2-5}alkenylR^{29}$ (wherein R²⁹ is as defined herein);

- 12) C₂₋₅alkynylR²⁹ (wherein R²⁹ is as defined herein);
- 13) C₁₋₅alkylX⁶R²⁹ (wherein X⁶ represents -O-, -S-, -SO-, -SO₂-, -NR³⁴C(O)-, -C(O)NR³⁵-, -SO₂NR³⁶-, -NR³⁷SO₂- or -NR³⁸- (wherein R³⁴, R³⁵, R³⁶, R³⁷ and R³⁸ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²⁹ is as defined herein);
- 14) C₂₋₅alkenylX⁷R²⁹ (wherein X⁷ represents -O-, -S-, -SO-, -SO₂-, -NR³⁹C(O)-, -C(O)NR⁴⁰-, -SO₂NR⁴¹-, -NR⁴²SO₂- or -NR⁴³- (wherein R³⁹, R⁴⁰, R⁴¹, R⁴² and R⁴³ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²⁹ is as defined herein);
- 15) C₂₋₅alkynylX⁸R²⁹ (wherein X⁸ represents -O-, -S-, -SO-, -SO₂-, -NR⁴⁴C(O)-, -C(O)NR⁴⁵-, -SO₂NR⁴⁶-, -NR⁴⁷SO₂- or -NR⁴⁸- (wherein R⁴⁴, R⁴⁵, R⁴⁶, R⁴⁷ and R⁴⁸ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²⁹ is as defined herein);
- 16) C₁₋₄alkylX⁹C₁₋₄alkylR²⁹ (wherein X⁹ represents -O-, -S-, -SO-, -SO₂-, -NR⁴⁹C(O)-, -C(O)NR⁵⁰-, -SO₂NR⁵¹-, -NR⁵²SO₂- or -NR⁵³- (wherein R⁴⁹, R⁵⁰, R⁵¹, R⁵² and R⁵³ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R²⁹ is as defined herein);
- 17) C₁₋₄alkylX⁹C₁₋₄alkylR²⁸ (wherein X⁹ and R²⁸ are as defined herein);
- 18) C₂₋₅alkenyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, C₁₋₄alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;
- 19) C₂₋₅alkynyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, C₁₋₄alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;
- 20) C₂₋₅alkenylX⁹C₁₋₄alkylR²⁸ (wherein X⁹ and R²⁸ are as defined herein);
- 21) C₂₋₅alkynylX⁹C₁₋₄alkylR²⁸ (wherein X⁹ and R²⁸ are as defined herein); and
- 22) C₁₋₄alkylR⁵⁴(C₁₋₄alkyl)_q(X⁹)_rR⁵⁵ (wherein X⁹ is as defined herein, q is 0 or 1, r is 0 or 1, and R⁵⁴ and R⁵⁵ are each independently selected from hydrogen, C₁₋₃alkyl, cyclopentyl, cyclohexyl and a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which C₁₋₃alkyl group may bear

1 or 2 substituents selected from oxo, hydroxy, halogeno and C₁₋₄alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₄cyanoalkyl, C₁₋₄alkyl, C₁₋₄hydroxyalkyl, C₁₋₄alkoxy, C₁₋₄alkoxyC₁₋₄alkyl, C₁₋₄alkylsulphonylC₁₋₄alkyl, C₁₋₄alkoxycarbonyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, C₁₋₄alkylaminoC₁₋₄alkoxy, di(C₁₋₄alkyl)aminoC₁₋₄alkoxy and a group -(O-)_f(C₁₋₄alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl), with the proviso that R⁵⁴ cannot be hydrogen);

and additionally wherein any C₁₋₅alkyl, C₂₋₅alkenyl or C₂₋₅alkynyl group in R⁵X¹- may bear one or more substituents selected from hydroxy, halogeno and amino;

R¹ represents hydrogen, oxo, halogeno, hydroxy, C₁₋₄alkoxy, C₁₋₄alkyl, C₁₋₄alkoxymethyl, C₁₋₄alkanoyl, C₁₋₄haloalkyl, cyano, amino, C₂₋₅alkenyl, C₂₋₅alkynyl, C₁₋₃alkanoyloxy, nitro, C₁₋₄alkanoylamino, C₁₋₄alkoxycarbonyl, C₁₋₄alkylsulphanyl, C₁₋₄alkylsulphinyl, C₁₋₄alkylsulphonyl, carbamoyl, N-C₁₋₄alkylcarbamoyl, N,N-di(C₁₋₄alkyl)carbamoyl, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl, N,N-di(C₁₋₄alkyl)aminosulphonyl, N-(C₁₋₄alkylsulphonyl)amino, N-(C₁₋₄alkylsulphonyl)-N-(C₁₋₄alkyl)amino, N,N-di(C₁₋₄alkylsulphonyl)amino, a C₃₋₇alkylene chain joined to two ring C carbon atoms, C₁₋₄alkanoylaminoC₁₋₄alkyl, carboxy,

or R¹ represents a group R⁵⁶X¹⁰, wherein X¹⁰ represents a direct bond, -O-, -CH₂-, -OC(O)-, -C(O)-, -S-, -SO-, -SO₂-, -NR⁵⁷C(O)-, -C(O)NR⁵⁸-, -SO₂NR⁵⁹-, -NR⁶⁰SO₂- or -NR⁶¹- (wherein R⁵⁷, R⁵⁸, R⁵⁹, R⁶⁰ and R⁶¹ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl), and R⁵⁶ is selected from one of the following twenty-two groups:

- 1) hydrogen, oxiranylC₁₋₄alkyl or C₁₋₅alkyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, chloro, bromo and amino;
- 2) C₁₋₅alkylX¹¹C(O)R⁶² (wherein X¹¹ represents -O- or -NR⁶³- (in which R⁶³ represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁶² represents C₁₋₃alkyl, -NR⁶⁴R⁶⁵ or

- OR⁶⁶ (wherein R⁶⁴, R⁶⁵ and R⁶⁶ which may be the same or different each represents hydrogen, C₁₋₅alkyl or C₁₋₃alkoxyC₂₋₃alkyl));
- 3) C₁₋₅alkylX¹²R⁶⁷ (wherein X¹² represents -O-, -S-, -SO-, -SO₂-, -OC(O)-, -NR⁶⁸C(O)-, -C(O)NR⁶⁹-, -SO₂NR⁷⁰-, -NR⁷¹SO₂- or -NR⁷²- (wherein R⁶⁸, R⁶⁹, R⁷⁰, R⁷¹ and R⁷² each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁶⁷ represents hydrogen, C₁₋₃alkyl, cyclopentyl, cyclohexyl or a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which C₁₋₃alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and C₁₋₄alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₄cyanoalkyl, C₁₋₄alkyl, C₁₋₄hydroxyalkyl, C₁₋₄alkoxy, C₁₋₄alkoxyC₁₋₄alkyl, C₁₋₄alkylsulphonylC₁₋₄alkyl, C₁₋₄alkoxycarbonyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, C₁₋₄alkylaminoC₁₋₄alkoxy, di(C₁₋₄alkyl)aminoC₁₋₄alkoxy and a group -(O-)_f(C₁₋₄alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl));
- 4) C₁₋₅alkylX¹³C₁₋₅alkylX¹⁴R⁷³ (wherein X¹³ and X¹⁴ which may be the same or different are each -O-, -S-, -SO-, -SO₂-, -NR⁷⁴C(O)-, -C(O)NR⁷⁵-, -SO₂NR⁷⁶-, -NR⁷⁷SO₂- or -NR⁷⁸- (wherein R⁷⁴, R⁷⁵, R⁷⁶, R⁷⁷ and R⁷⁸ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁷³ represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl);
- 5) R⁷⁹ (wherein R⁷⁹ is a 4-6-membered saturated heterocyclic group (linked via carbon or nitrogen) with 1-2 heteroatoms, selected independently from O, S and N, which heterocyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₄cyanoalkyl, C₁₋₄alkyl, C₁₋₄hydroxyalkyl, C₁₋₄alkoxy, C₁₋₄alkoxyC₁₋₄alkyl, C₁₋₄alkylsulphonylC₁₋₄alkyl, C₁₋₄alkoxycarbonyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, C₁₋₄alkylaminoC₁₋₄alkoxy, di(C₁₋₄alkyl)aminoC₁₋₄alkoxy and a group -(O-)_f(C₁₋₄alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered

- saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl));
- 6) C₁₋₅alkylR⁷⁹ (wherein R⁷⁹ is as defined herein);
- 7) C₂₋₅alkenylR⁷⁹ (wherein R⁷⁹ is as defined herein);
- 8) C₂₋₅alkynylR⁷⁹ (wherein R⁷⁹ is as defined herein);
- 9) R⁸⁰ (wherein R⁸⁰ represents a pyridone group, a phenyl group or a 5-6-membered aromatic heterocyclic group (linked via carbon or nitrogen) with 1-3 heteroatoms selected from O, N and S, which pyridone, phenyl or aromatic heterocyclic group may carry up to 5 substituents selected from hydroxy, halogeno, amino, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄hydroxyalkyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, C₁₋₄hydroxyalkoxy, carboxy, trifluoromethyl, cyano, -C(O)NR⁸¹R⁸², -NR⁸³C(O)R⁸⁴ (wherein R⁸¹, R⁸², R⁸³ and R⁸⁴, which may be the same or different, each represents hydrogen, C₁₋₄alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and a group -(O-)_f(C₁₋₄alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl));
- 10) C₁₋₅alkylR⁸⁰ (wherein R⁸⁰ is as defined herein);
- 11) C₂₋₅alkenylR⁸⁰ (wherein R⁸⁰ is as defined herein);
- 12) C₂₋₅alkynylR⁸⁰ (wherein R⁸⁰ is as defined herein);
- 13) C₁₋₅alkylX¹⁵R⁸⁰ (wherein X¹⁵ represents -O-, -S-, -SO-, -SO₂-, -NR⁸⁵C(O)-, -C(O)NR⁸⁶-, -SO₂NR⁸⁷-, -NR⁸⁸SO₂- or -NR⁸⁹- (wherein R⁸⁵, R⁸⁶, R⁸⁷, R⁸⁸ and R⁸⁹ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁸⁰ is as defined herein);
- 14) C₂₋₅alkenylX¹⁶R⁸⁰ (wherein X¹⁶ represents -O-, -S-, -SO-, -SO₂-, -NR⁹⁰C(O)-, -C(O)NR⁹¹-, -SO₂NR⁹²-, -NR⁹³SO₂- or -NR⁹⁴- (wherein R⁹⁰, R⁹¹, R⁹², R⁹³ and R⁹⁴ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁸⁰ is as defined herein);
- 15) C₂₋₅alkynylX¹⁷R⁸⁰ (wherein X¹⁷ represents -O-, -S-, -SO-, -SO₂-, -NR⁹⁵C(O)-, -C(O)NR⁹⁶-, -SO₂NR⁹⁷-, -NR⁹⁸SO₂- or -NR⁹⁹- (wherein R⁹⁵, R⁹⁶, R⁹⁷, R⁹⁸ and R⁹⁹ each

independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁸⁰ is as defined herein);

- 16) C₁₋₄alkylX¹⁸C₁₋₄alkylR⁸⁰ (wherein X¹⁸ represents -O-, -S-, -SO-, -SO₂-, -NR¹⁰⁰C(O)-, -C(O)NR¹⁰¹-, -SO₂NR¹⁰²-, -NR¹⁰³SO₂- or -NR¹⁰⁴- (wherein R¹⁰⁰, R¹⁰¹, R¹⁰², R¹⁰³ and R¹⁰⁴ each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl) and R⁸⁰ is as defined herein);
- 17) C₁₋₄alkylX¹⁸C₁₋₄alkylR⁷⁹ (wherein X¹⁸ and R⁷⁹ are as defined herein);
- 18) C₂₋₅alkenyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, C₁₋₄alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;
- 19) C₂₋₅alkynyl which may be unsubstituted or which may be substituted with one or more groups selected from hydroxy, fluoro, amino, C₁₋₄alkylamino, N,N-di(C₁₋₄alkyl)amino, aminosulphonyl, N-C₁₋₄alkylaminosulphonyl and N,N-di(C₁₋₄alkyl)aminosulphonyl;
- 20) C₂₋₅alkenylX¹⁸C₁₋₄alkylR⁷⁹ (wherein X¹⁸ and R⁷⁹ are as defined herein);
- 21) C₂₋₅alkynylX¹⁸C₁₋₄alkylR⁷⁹ (wherein X¹⁸ and R⁷⁹ are as defined herein); and
- 22) C₁₋₄alkylR¹⁰⁵(C₁₋₄alkyl)_x(X¹⁸)_yR¹⁰⁶ (wherein X¹⁸ is as defined herein, x is 0 or 1, y is 0 or 1, and R¹⁰⁵ and R¹⁰⁶ are each independently selected from hydrogen, C₁₋₃alkyl, cyclopentyl, cyclohexyl and a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which C₁₋₃alkyl group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno and C₁₋₄alkoxy and which cyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₄cyanoalkyl, C₁₋₄alkyl, C₁₋₄hydroxyalkyl, C₁₋₄alkoxy, C₁₋₄alkoxyC₁₋₄alkyl, C₁₋₄alkylsulphonylC₁₋₄alkyl, C₁₋₄alkoxycarbonyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, C₁₋₄alkylaminoC₁₋₄alkoxy, di(C₁₋₄alkyl)aminoC₁₋₄alkoxy and a group -(-O-)_f(C₁₋₄alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl) with the proviso that R¹⁰⁵ cannot be hydrogen);

and additionally wherein any C₁₋₅alkyl, C₂₋₅alkenyl or C₂₋₅alkynyl group in R⁵⁶X¹⁰ - may bear one or more substituents selected from hydroxy, halogeno and amino;

R^{2a} represents hydrogen, halogeno, C₁₋₃alkyl, trifluoromethyl, C₁₋₃alkoxy, C₁₋₃alkylsulphanyl, -NR^{3a}R^{4a} (wherein R^{3a} and R^{4a}, which may be the same or different, each represents hydrogen or C₁₋₃alkyl), or R^{5a}(CH₂)_{za}X^{1a} (wherein R^{5a} is a 4-, 5- or 6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which heterocyclic group may bear 1 or 2 substituents selected from oxo, hydroxy, halogeno, cyano, C₁₋₄cyanoalkyl, C₁₋₄alkyl, C₁₋₄hydroxyalkyl, C₁₋₄alkoxy, C₁₋₄alkoxyC₁₋₄alkyl, C₁₋₄alkylsulphonylC₁₋₄alkyl, C₁₋₄alkoxycarbonyl, C₁₋₄aminoalkyl, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkylaminoC₁₋₄alkyl, di(C₁₋₄alkyl)aminoC₁₋₄alkyl, C₁₋₄alkylaminoC₁₋₄alkoxy, di(C₁₋₄alkyl)aminoC₁₋₄alkoxy and a group -(O-)_f(C₁₋₄alkyl)_gringD (wherein f is 0 or 1, g is 0 or 1 and ring D is a 4-6-membered saturated heterocyclic group with 1-2 heteroatoms, selected independently from O, S and N, which cyclic group may bear one or more substituents selected from C₁₋₄alkyl), za is an integer from 0 to 4 and X^{1a} represents a direct bond, -O-, -CH₂-, -S-, -SO-, -SO₂-, -NR^{6a}C(O)-, -C(O)NR^{7a}-, -SO₂NR^{8a}-, -NR^{9a}SO₂- or -NR^{10a}- (wherein R^{6a}, R^{7a}, R^{8a}, R^{9a} and R^{10a} each independently represents hydrogen, C₁₋₃alkyl or C₁₋₃alkoxyC₂₋₃alkyl));

or a salt or prodrug thereof;

with the proviso that R² is not hydrogen, and excluding the compounds:

6,7-dimethoxy-4-(1-naphthylsulphanyl)quinazoline,

6,7-dimethoxy-4-(2-naphthylsulphanyl)quinazoline,

6,7-dimethoxy-4-(1-naphthyloxy)quinazoline and

6,7-dimethoxy-4-(2-naphthyloxy)quinazoline.